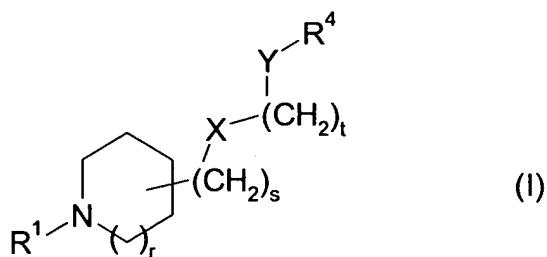


CLAIMS

1. A compound of the general formula (I):



wherein

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R¹ is

C₁₋₆-alkyl, C₂₋₆-alkenyl, C₂₋₆-alkynyl,

- which may optionally be substituted with one or more substituents independently selected from R¹¹, wherein R¹¹ is halogen, C₁₋₆-alkoxy or hydroxy,

C₃₋₈-cycloalkyl, C₅₋₈-cycloalkenyl, C₃₋₈-cycloalkyl-C₁₋₆-alkyl, di(C₃₋₈-cycloalkyl)-C₁₋₆-alkyl,

C₃₋₈-cycloalkyl-C₂₋₆-alkenyl, C₃₋₈-cycloalkyl-C₂₋₆-alkynyl, C₅₋₈-cycloalkenyl-C₁₋₆-alkyl,

C₅₋₈-cycloalkenyl-C₂₋₆-alkenyl, C₅₋₈-cycloalkenyl-C₂₋₆-alkynyl, 4-pyridyl or tetrahydropyranyl,

- 15 • wherein the cyclic moieties may optionally be substituted with one or more substituents independently selected from R¹², wherein R¹² is C₁₋₆-alkyl, halogen, trifluoromethyl or 2,2,2-trifluoroethyl,

r is 0, 1 or 2,

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s is 0, 1, 2 or 3,

t is 0, 1, 2 or 3,

- 25 X is C=O, CHOH or CR²R³; wherein R² and R³ independently are hydrogen or C₁₋₆-alkyl, or X is a bond,

Y is heteroaryl optionally substituted with one or more substituents independently selected from R¹⁸,

R¹⁸ is halogen, nitro, cyano, hydroxy, C₁₋₆-alkyl, C₁₋₆-alkylthio or C₁₋₆-alkoxy.

R⁴ is

- 5 (a) C₁₋₆-alkyl, C₃₋₈-cycloalkyl or C₃₋₈-cycloalkenyl, which may optionally be substituted with one or more substituents independently selected from R¹³, wherein R¹³ is C₃₋₈-cycloalkyl, C₁₋₆-alkoxy, C₁₋₆-alkylthio, cyano, halo-C₁₋₆-alkyl, halo-C₁₋₆-alkoxy, and halogen,

or

- 10 (b) aryl, aryl-C₁₋₆-alkyl, aryl-C₂₋₆-alkenyl, or heteroaryl

which may optionally be substituted with one or more substituents independently selected from R¹⁴

R¹⁴ is

- 15 ▪ halogen, nitro, cyano, acyl, hydroxy, C₁₋₆-alkyl, C₁₋₆-alkylthio, C₁₋₆-alkylsulfonyl, C₁₋₆-alkylsulfonyloxy, C₁₋₆-alkoxy, C₃₋₈-cycloalkyl, halo-C₁₋₆-alkyl, halo-C₁₋₆-alkoxy, -NR⁵R⁶, R⁵R⁶N-C₁₋₆-alkyl-, R⁵R⁶N-C₁₋₆-alkoxy-, or -O(C=O)NR⁵R⁶, or wherein two substituents in adjacent positions together form a radical -O-(CH₂)₁₋₃-O-, wherein R⁵ and R⁶ independently are hydrogen, C₁₋₆-alkyl, C₃₋₈-cycloalkyl, C₁₋₆-alkanoyl or aryl, or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 3 to 7 membered, saturated or unsaturated ring, which may be fused to a benzene ring,
- 20 ▪ a group of the formula

- 25 -(W)_k-A wherein

W is -C₁₋₆-alkyl-, -(O), -C₂₋₆-alkenyl-, -(O)-C₁₋₆-alkyl-O-, -(CH₂)_n-(C=O)-(CH₂)_m-, -O-

wherein

- 30 I is 0 or 1
k is 0 or 1
n and m are independently 0, 1, 2 or 3,
A is

- o aryl, aryl-C₁₋₆-alkyl, heteroaryl, heteroaryl-C₁₋₆-alkyl, C₁₋₆-alkyl or C₃₋₈-cycloalkyl wherein the ring moieties optionally may be substituted with one or more substituents independently selected from R¹⁵

R¹⁵ is

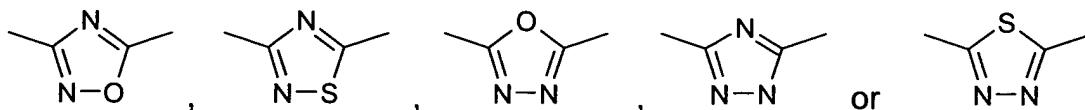
- 5
 - halogen, nitro, cyano, hydroxy, C₁₋₆-alkylthio, C₁₋₆-alkylsulfonyl, C₁₋₆-alkylsulfonyloxy, C₁₋₆-alkyl, C₁₋₆-alkoxy, C₃₋₈-cycloalkyl, halo-C₁₋₆-alkyl, halo-C₁₋₆-alkoxy, -NR⁷R⁸, R⁷R⁸N-C₁₋₆-alkyl-, R⁷R⁸N-C₁₋₆-alkoxy-, or -O(C=O)NR⁷R⁸, or wherein two substituents in adjacent positions together form a radical -O-(CH₂)₁₋₃-O-,
- 10 wherein R⁷ and R⁸ independently are hydrogen, C₁₋₆-alkyl, C₃₋₈-cycloalkyl, C₁₋₆-alkanoyl or aryl, or R⁷ and R⁸ together with the nitrogen atom to which they are attached form a 3 to 7 membered, saturated or unsaturated ring, which may be fused to a benzene ring,
- 15
 - o NR⁹R¹⁰ wherein R⁹ and R¹⁰ independently are hydrogen, C₁₋₆-alkyl, C₃₋₈-cycloalkyl, C₁₋₆-alkanoyl or aryl, or R⁹ and R¹⁰ together with the nitrogen atom to which they are attached form a 3 to 7 membered, saturated or unsaturated ring, which may be fused to a benzene ring, and the ring may contain further heteroatoms and it may optionally be substituted with one or more substituents independently selected from R¹⁶,
- 20 wherein R¹⁶ is C₁₋₆-alkyl, C₃₋₈-cycloalkyl, C₁₋₆-alkanoyl or aryl optionally substituted with one or more substituents independently selected from R¹⁷, wherein R¹⁷ is halogen, nitro, cyano, hydroxy, or C₁₋₆-alkyl;

as well as any diastereomer or enantiomer or tautomeric form, mixtures of these, or a pharmaceutically acceptable salt thereof.

2. A compound according to claim 1, wherein R¹ is C₃₋₈-cycloalkyl or C₁₋₆-alkyl.
3. A compound according to claim 2 wherein R¹ is cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, methyl, ethyl, propyl, 1-methylpropyl, 1-ethyl-propyl, isopropyl, or *tert*-butyl.
4. A compound according to claim 3 wherein R¹ is cyclopropyl, cyclopentyl, 1-ethyl-propyl, or isopropyl.
- 35 5. A compound according to claim 4 wherein R¹ is isopropyl.

6. A compound according to claim 4 wherein R¹ is cyclopropyl.
7. A compound according to claim 1, wherein X is a bond.
- 5 8. A compound according to claim 1, wherein s and t together are 0, 1, 2 or 3.
9. A compound according to claim 1 wherein r is 1.
10. A compound according to claim 1 wherein s is 0 or 1.
- 10 11. A compound according to claim 10 wherein s is 0.
12. A compound according to claim 1 wherein t is 0.
- 15 13. A compound according to claim 1, wherein Y is a 5- or 6-membered heterocyclic aromatic ring system optionally substituted with one or more substituents independently selected from R¹⁸.
14. A compound according to claim 13, wherein Y is a 5-membered heterocyclic aromatic ring system optionally substituted with one or more substituents independently selected from R¹⁸.
- 20 15. A compound according to claim 14, wherein Y is a 5-membered heterocyclic aromatic ring system containing 1, 2 or 3 heteroatoms, optionally substituted with one or more substituents independently selected from R¹⁸.
- 25 16. A compound according to claim 15, wherein Y is a 5-membered heterocyclic aromatic system containing 3 heteroatoms, optionally substituted with one or more substituents independently selected from R¹⁸.
- 30 17. A compound according to claim 16, wherein Y is selected from the group consisting of oxadiazolyl, thiadiazolyl, or triazolyl, optionally substituted with one or more substituents independently selected from R¹⁸.
- 35 18. A compound according to claim 17, wherein Y is selected from

91



19. A compound according to claim 1, wherein R⁴ is aryl, aryl-C₁₋₆-alkyl, either of which may optionally be substituted with one or more substituents independently selected from R¹⁴, or
5 C₃₋₈-cycloalkyl optionally substituted with one or more substituents independently selected from R¹³.
20. A compound according to claim 19 wherein R⁴ is aryl optionally substituted with one or more substituents independently selected from R¹⁴.
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21. A compound according to claim 19 wherein R⁴ is phenyl, biphenyl, or naphthyl optionally substituted with one or more substituents independently selected from R¹⁴.
22. A compound according to claim 21, wherein R⁴ is phenyl optionally substituted with one
15 or more substituents independently selected from R¹⁴.
23. A compound according to claim 1 wherein R¹³ is C₁₋₆-alkyl.
24. A compound according to claim 1 wherein R¹⁴ is
20 halogen, cyano, hydroxy, C₁₋₆-alkyl, C₁₋₆-alkylsulfonyl, C₁₋₆-alkylsulfonyloxy,
C₁₋₆-alkoxy, C₃₋₈-cycloalkyl, -CF₃, -OCF₃, -NR⁵R⁶, R⁵R⁶N-C₁₋₆-alkyl-, or
a group of the formula -(W)_k-A.
25. A compound according to claim 24 wherein R¹⁴ is
25 F, Cl, cyano, methyl, ethyl, propyl, butyl, tert-butyl, methyl-sulfonyl, methylsulfonyloxy,
methoxy, ethoxy, cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, -CF₃, -OCF₃, -
NR⁵R⁶, R⁵R⁶N-methyl-, or
a group of the formula -(W)_k-A.
- 30 26. A compound according to claim 25 wherein R¹⁴ is
F, Cl, cyano, methyl, tert-butyl, methyl-sulfonyl, methoxy, cyclopentyl, cyclohexyl,
-CF₃, -OCF₃, -NR⁵R⁶, R⁵R⁶N-methyl-, or
a group of the formula -(W)_k-A.

27. A compound according to claim 26 wherein R¹⁴ is a group of the formula -(W)_k-A.
28. A compound according to claim 1, wherein k is 1.
- 5 29. A compound according to claim 1 wherein k is 0.
30. A compound according to claim 1 wherein W is -C₁₋₆-alkyl-, -(O)-C₁₋₆-alkyl-O-, -(CH₂)_n-(C=O)-(CH₂)_m-, or -O-.
- 10 31. A compound according to claim 30 wherein W is -C₁₋₆-alkyl- or -(CH₂)_n-(C=O)-(CH₂)_m-.
32. A compound according to claim 31 wherein W is methylene, ethylene, propylene or -(CH₂)_n-(C=O)-(CH₂)_m-.
- 15 33. A compound according to claim 1 wherein n is 0 or 1.
34. A compound according to claim 33 wherein n is 0.
35. A compound according to any one of the claims 1 to 34 wherein m is 0 or 1.
- 20 36. A compound according to claim 35 wherein m is 0.
37. A compound according to claim 1 wherein l is 0.
- 25 38. A compound according to claim 1 wherein A is C₁₋₆-alkyl, aryl or C₃₋₈-cycloalkyl, wherein the ring moieties optionally may be substituted with one or more substituents independently selected from R¹⁵, or A is NR⁹R¹⁰.
39. A compound according to claim 38 wherein A is methyl, ethyl, phenyl, cyclopropyl,
- 30 40. A compound according to claim 39 wherein A is phenyl optionally substituted with one or more substituents independently selected from R¹⁵, or A is NR⁹R¹⁰.
40. A compound according to claim 39 wherein A is phenyl optionally substituted with one or more substituents independently selected from R¹⁵.

41. A compound according to claim 40 wherein A is phenyl.

42. A compound according to claim 39 wherein A is NR⁹R¹⁰.

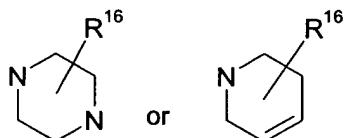
5 43. A compound according to claim 1 wherein R¹⁵ is halogen, nitro, cyano, hydroxy, C₁₋₆-alkylthio, C₁₋₆-alkylsulfonyl, C₁₋₆-alkylsulfonyloxy, C₁₋₆-alkyl, C₁₋₆-alkoxy, C₃₋₈-cycloalkyl, halo-C₁₋₆-alkyl, or halo-C₁₋₆-alkoxy.

10 44. A compound according to claim 43 wherein R¹⁵ is halogen, cyano, hydroxy, CH₃-S-, CH₃CH₂-S-, methylsulfonyl, methylsulfonyloxy, methyl, ethyl, propyl, butyl, isopropyl, methoxy, ethoxy, cyclopropyl, cyclobutyl, cyclopentyl, or cyclohexyl, -CF₃, or -OCF₃.

45. A compound according to claim 44 wherein R¹⁵ is halogen, methyl, ethyl, methoxy, ethoxy, -CF₃, or -OCF₃.

15 46. A compound according to claim 1 wherein R⁹ and R¹⁰ together with the nitrogen atom to which they are attached form a 3 to 7 membered, saturated or unsaturated ring, which may be fused to a benzene ring, and the ring may contain further heteroatoms and it may optionally be substituted with one or more substituents independently selected from R¹⁶.

20 47. A compound according to claim 46 wherein R⁹ and R¹⁰ together with the nitrogen atom to which they are attached form a structure selected from



25 48. A compound according to claim 1 wherein R¹⁶ is methyl, ethyl, 1-ethyl-propyl or phenyl optionally substituted with one or more substituents independently selected from R¹⁷.

49. A compound according to claim 1 wherein R¹⁷ is halogen.

30 50. A pharmaceutical composition comprising, as an active ingredient, at least one compound according to claim 1 together with one or more pharmaceutically acceptable carriers or excipients.

51. A pharmaceutical composition according to claim 50 in unit dosage form, comprising from about 0.05 mg to about 1000 mg, preferably from about 0.1 mg to about 500 mg and especially preferred from about 0.5 mg to about 200 mg of the compound.

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52. A pharmaceutical composition according to claim 50 wherein the compound exhibits histamine H3 antagonistic activity or histamine H3 inverse agonistic activity.

10 53. A method for treating diseases and disorders in which an inhibition of the H3 histamine receptor has a beneficial effect comprising administering to a subject in need thereof a therapeutically effective amount of a compound according to claim 1.

54. A method for the reduction of body weight, comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

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55. A method for the treatment of overweight or obesity, comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

20 56. A method for the suppression of appetite or for satiety induction comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

57. A method for the prevention and/or treatment of disorders and diseases related to overweight or obesity comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

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58. A method for the prevention and/or treatment of eating disorders such as bulimia and binge eating comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

30 59. A method for the treatment of IGT comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

60. A method for the treatment of type 2 diabetes comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1

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61. A method for the delaying or prevention of the progression from IGT to type 2 diabetes comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

5 62. A method for the delaying or prevention of the progression from non-insulin requiring type 2 diabetes to insulin requiring type 2 diabetes comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1

10 63. A method for the treatment of diseases and disorders in which a stimulation of the H3 histamine receptor has a beneficial effect comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

64. A compound according to Claim 1 exhibiting histamine H3 agonistic activity.

15 65. A method of treating allergic rhinitis, ulcer or anorexia comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

20 66. A method for the treatment of Alzheimer's disease, narcolepsy or attention deficit disorder comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

67. The method according to claim 53 wherein the therapeutically effective amount of the compound is in the range of from about 0.05 mg to about 2000 mg, preferably from about 0.1 mg to about 1000 mg and especially preferred from about 0.5 mg to about 500 mg per day.